



From the INTERNATIONAL BUREAU

PCT

NOTIFICATION OF ELECTION

(PCT Rule 61.2)

To:

Assistant Commissioner for Patents United States Patent and Trademark Office Box PCT Washington, D.C.20231 ÉTATS-UNIS D'AMÉRIQUE

Date of mailing (day/month/year) 14 December 1999 (14.12.99)	in its capacity as elected Office
International application No. PCT/US99/08802	Applicant's or agent's file reference 02441.80414
International filing date (day/month/year) 22 April 1999 (22.04.99)	Priority date (day/month/year) 22 April 1998 (22.04.98)
Applicant	
PALIARD, Xavier	

L		PALIARD, Xavier
	1.	The designated Office is hereby notified of its election made:
		X in the demand filed with the International Preliminary Examining Authority on:
		19 November 1999 (19.11.99)
		in a notice effecting later election filed with the International Bureau on:
	2.	The election X was
		was not
		made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b).
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The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland

Authorized officer

A. Karkachi

Telephone No.: (41-22) 338.83.38

Facsimile No.: (41-22) 740.14.35

From the:

INTERNATIONAL PRELIMINARY EXAMINING AUTHORITY

To:

BLACKBURN, Robert P. CHIRON CORPORATION Intellectual Property - R338 P.O. Box 8097

Emeryville, CA 94662-8097 ETATS-UNIS D'AMERIQUE



WRITTEN OPINION

(PCT Rule 66)

Date of mailing (day/month/year)	31.03.2000	
REPLY DUE	within 3 month(s) from the above date of mailing	

Applicant's or agent's file reference

International application No.

PCT/US99/08802

1521.100

International filing date (day/month/year)

22/04/1999

Priority date (day/month/year)

22/04/1998

International Patent Classification (IPC) or both national classification and IPC

C12N15/36

Applicant

CHIRON CORPORATION et al.

- 1. This written opinion is the first drawn up by this International Preliminary Examining Authority.
- 2. This opinion contains indications relating to the following items:
 - ☑ Basis of the opinion
 - II Priority
 - III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability
 - IV 🛛 Lack of unity of invention
 - V Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement
 - VI

 Certain document cited
 - VII Certain defects in the international application
 - VIII

 Certain observations on the international application
- 3. The applicant is hereby invited to reply to this opinion.

When? See the time limit indicated above. The applicant may, before the expiration of that time limit,

request this Authority to grant an extension, see Rule 66.2(d).

How? By submitting a written reply, accompanied, where appropriate, by amendments, according to Rule 66.3.

For the form and the language of the amendments, see Rules 66.8 and 66.9.

Also:

For an additional opportunity to submit amendments, see Rule 66.4.

For the examiner's obligation to consider amendments and/or arguments, see Rule 66.4 bis.

For an informal communication with the examiner, see Rule 66.6.

If no reply is filed, the international preliminary examination report will be established on the basis of this opinion.

4. The final date by which the international preliminary examination report must be established according to Rule 69.2 is: 22/08/2000.

AAH FA 413/09/00

Authorized officer / Examiner 1999

Name and mailing address of the international preliminary examining authority:

9)

European Patent Office D-80298 Munich

Tel. +49 89 2399 - 0 Tx: 523656 epmu d

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Petri, B

Formalities officer (incl. extension of time limits)

Vullo, C

Telephone No. +49 89 2399 8061



WRITTEN OPINION

 Basis of the opi 	inion
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••		no or the opinion			
1.	This	s opinion has been esponse to an invita	drawn on the basis of (su ation under Article 14 are	ubstitute sheets which have been furnished to the re referred to in this opinion as "originally filed".):	eceiving Office
	Des	cription, pages:			
	1-14	1	as originally filed		
	Cla	ims, No.:			
	1-29	9	as originally filed		-
	Dra	wings, sheets:			
	1/5-	5/5	as originally filed		
2.	The	amendments have	e resulted in the cancellati	tion of:	
		the description,	pages:	,e	
		the claims,	Nos.:	r ·	
		the drawings,	sheets:	(
3.	This	s opinion has been sidered to go beyo	established as if (some o	of) the amendments had not been made, since they (Rule 70.2(c)):	have been
			ا		
4.	Add	litional observation	s, if necessary:		
				<u> </u>	
			·		
IV	. Lac	k of unity of inve	ntion -		
1.	In r	esponse to the invit	tation (Form PCT/IPEA/40	05) to restrict or pay additional fees, the applicant h	as:
		restricted the clair	ns.		
		paid additional fee	es.		
		paid additional fee	es under protest.		
		neither restricted	nor paid additional fees.		

2.

This Authority found that the requirement of unity of invention is not complied with for the following reasons



and chose, according to Rule 68.1, not to invite the applicant to restrict or pay additional fees:

see separate sheet

3.	Consequently, the following parts of the international application were the subject of international preliminary
	examination in establishing this opinion:

☑ all parts.

☐ the parts relating to claims Nos. .

V. Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N)

Claims 1-2, 5-8, 10-21, 25-29

Inventive step (IS)

Claims 3-4, 23-24

Industrial applicability (IA)

Claims

2. Citations and explanations

see separate sheet

VIII. Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made:

see separate sheet



Re Item IV

Lack of unity of invention

The international search report has been drawn up in respect of the entire international application but the IPEA is of the opinion that the application does not comply with the requirements of unity of invention as set forth in the PCT regulations (Article 34(3), Rules 13 and 68 PCT):

The separate inventions/groups of inventions are:

An immunogenic composition, comprising a DNA immunogen; and a chemokine or a polynucleotide encoding a chemokine and methods of enhancing an immune response comprising administering a chemokine and a DNA immunogen, wherein the immunogen is:

- (i) a viral immunogen and the chemokine is MIP-1a
- (ii) an immunogen expressed by a tumor and the chemokine is MIP-1a
- (iii) a viral immunogen and the chemokine is BLC
- (iv) an immunogen expressed by a tumor and the chemokine is BLC

The common concept linking together above group of inventions is the immunogenic composition, comprising a DNA immunogen; and a chemokine or a polynucleotide encoding a chemokine (claim 1).

This concept is not novel (see item V).

Furthermore as also viral immunogenes are known in the composition of claim 1, additional group of inventions arise, consisting of hepatitis C virus non-structural polypeptides (claim 3) and HIV polypeptides (claim 5) and any subject-matter resulting from combinations with the chemokines of claim 8 and 9.

However, in the light of the findings below (see item V) and since all the invention groups can be subjected to IPE with reasonable additional effort, the IPER will cover all group of inventions.

The applicant must be aware of a possible objection because of lack of unity in any



regional phase of the present application.

Re Item V

Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

Reference is made to the following documents:

- D1: WO 96 36366 A (DOW STEVE W ;ELMSLIE ROBYN E (US); NAT JEWISH CENTER FOR IMMUNOLO) 21 November 1996 (1996-11-21)
- D2: WO 96 11279 A (US HEALTH) 18 April 1996 (1996-04-18)
- D3: WO 98 15285 A (WISTAR INST ;ERTL HILDEGUND C J (US); THURIN MAGDALENA (US)) 16 April 1998 (1998-04-16)
- D4: KHUDYAKOV Y E ET AL: 'LINEAR B-CELL EPITOPES OF THE NS3-NS4-NS5 PROTEINS OF THE HEPATITISC VIRUS AS MODELED WITH SYNTHETIC PEPTIDES' VIROLOGY, vol. 206, 1 January 1995 (1995-01-01), pages 666-672, XP000574456 ISSN: 0042-6822

Novelty, Article 33(2), PCT

D1 discloses immunogenic compositions and methods of enhancing an immune response, comprising DNA immunogens coding for viral antigens and a chemokine or a polynucleotide encoding said chemokine (i.e. MIP1a) (page 6, line 23 - page 7 line 28; page 14 line 13; Examples 12-16; Claims 1, 2, 10-12, claim 53-62, claim 80-85). This disclosure anticipates the subject-matter of claims 1-2, 8, 10, 11-21, 27-29.

D2 discloses immunogenic compositions and methods of enhancing an immune response, comprising DNA immunogens coding for viral antigens (i.e. HIV, p17; page 27 line 14) and/or tumor antigens and a chemokine or a polynucleotide encoding said chemokine (i.e. MIP1a) (page 8, line 14-20; page 28, line 6; Claims 8-10; claim 18, claim 24-25). This disclosure anticipates the subject-matter of claims 1-2, 5-8, 10-21, 25-29.

WRITTEN OPINION SEPARATE SHEET

D3 discloses immunogenic compositions and methods of enhancing an immune response, comprising DNA immunogens coding for tumor antigens and a chemokine or a polynucleotide encoding said chemokine (page 4, line 24 - page 5 line 1; page 16 line 22-24; Example 13). This disclosure anticipates the subject-matter of claims 1-2, 7, 10, 11-20, 27-29.

In summary, in view of above prior art the subject-matter of claim 1-2, 5-8, 10-21, 25-29 is considered to lack novelty.

Inventive step, Article 33(3), PCT

Claim 3-4, 23-24 refer to HCV non structural polypeptides (i.e. NS3, NS4, NS5a,b) to be chosen as DNA immunogen. As however, these peptides are known to comprise strong antigenic epitopes recognized by antisera from patients infected with HCV (D4, Abstract, Table 1 and 2) their selection for the composition and methods of claim 1 and 11 is considered not to involve an inventive step.

As shown by the applicant, not all chemokines are equally suitable for the composition and method proposed in claim 1 and 11 (see below item VIII). As such, the selection of specific chemokines, that exhibit an enhancing effect on the response against a DNA immunogen is considered an inventive contribution to the art. Consequently employment of the chemokine BLC (claim 9, 22) in the composition of claim 1 and in the method of claim 11 appears to be inventive.

Industrial applicability, Article 33(4), PCT

For the assessment of the present claims 11-29 on the question whether they are industrially applicable, no unified criteria exist in the PCT Contracting States. The patentability can also be dependent upon the formulation of the claims. The EPO, for example, does not recognize as industrially applicable the subject-matter of claims to the use of a compound in medical treatment, but may allow, however, claims to a known compound for first use in medical treatment and the use of such a compound for the manufacture of a medicament for a new medical treatment.



WRITTEN OPINION SEPARATE SHEET

International application No. PCT/US99/08802

Re Item VI Certain documents cited

Certain published documents (Rule 70.10)

Application No

Publication date (day/month/year)

Filing date (day/month/year)

Priority date (valid claim) (day/month/year)

WO 99/29728

17.06.1999

11.12.1998

11.12.1997

Re Item VIII

Certain observations on the international application

The present application provides evidence that <u>selected</u> chemokines can enhance the immune response to a DNA immunogen.

Examples are provided which relate to the effect on cytoxic T cell response to HCV NS3 in a cytotoxic T cell assay after coadministration of plasmids encoding for specific HCV antigens (i.e. Δ NS3, NS4, NS5a and NS5b) and specific cytokines (GM-CSF) and/or chemokines (RANTES, MIP1 α) to baboons (Example 1). In addition, the effect of coadministration of HIV p55 with the chemokine BLC on anti-gag antibody titer is reported (Fig. 5).

Of the examples studied only MIP1 α and BLC have shown a positive effect on the immune response against the DNA immunogens (Table 1 and 2; Fig. 5). Consequently, as apparently <u>not all chemokines</u> promote an enhanced immune response, any broadening beyond the chemokines exemplified (i.e. MIP1 α claim 8, 21 and BLC claim 9, 22) is considered unjustified. Therefore, as also chemokines are embraced by the subject-matter of claims 1-7, 10, 11-20, 23-29 for which no positive effect has been shown (e.g. RANTES, Table 1), the invention cannot be carried out over the entire scope of said claims. Consequently claims 1-7, 10, 11-20, 23-29 are considered not to comply with the requirements of Article 5 and 6, PCT.



(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference	FOR FURTHER see Notification of Transmittal of International Search Report			
02441.80414	ACTION (Form PC1/ISA/	220) as well as, where applicable, item 5 below.		
International application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)		
PCT/US 99/08802	22/04/1999	22/04/1998		
Applicant				
CHIDON CORROBATION I				
CHIRON CORPORATION et al.				
This International Search Report has been	prepared by this International Searching Aut	booth, and in transmitted to the small and		
according to Article 18. A copy is being tra	nsmitted to the International Bureau.	anony and is transmitted to the applicant		
This International Search Report consists of	of a total of 4 sheets.			
	a copy of each prior art document cited in this	s report.		
Basis of the report				
a. With regard to the language, the in	nternational search was carried out on the ba	sis of the international application in the		
language in which it was filed, unle	ss otherwise indicated under this item.	olo of the international application in the		
the international search wa Authority (Rule 23.1(b)).	as carried out on the basis of a translation of t	he international application furnished to this		
b. With regard to any nucleotide and	Vor amino acid sequence disclosed in the in	nternational application, the international search		
was carried out on the basis of the contained in the internation	sequence listing : lial application in written form.			
filed together with the inter	national application in computer readable form	n.		
furnished subsequently to this Authority in written form.				
furnished subsequently to this Authority in computer readble form.				
the statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.				
the statement that the infor furnished	mation recorded in computer readable form is	s identical to the written sequence listing has been		
2. X Certain claims were found	d unsearchable (See Box I).			
3. Unity of invention is lacki	ng (see Box II).			
4. With regard to the title,				
the text is approved as sub	mitted by the applicant.			
	ed by this Authority to read as follows:			
ENHANCING IMMUNE RESPO	NSES TO GENETIC IMMUNIZATION	ON BY USING A CHEMOKINE		
5. With regard to the abstract,				
X the text is approved as sub				
the text has been established within one month from the control of	ed, according to Rule 38.2(b), by this Authorit late of mailing of this international search rep	y as it appears in Box III. The applicant may, ort, submit comments to this Authority.		
6. The figure of the drawings to be publish				
as suggested by the applica	•	X None of the figures.		
because the applicant failed				
because this figure better ch	naracterizes the invention.			

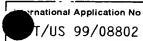


nternational application No.

PCT/US 99/08802

Box I	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Inte	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely: Remark: Although claims 11-29 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2.	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Intel	rnational Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4 r	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark o	The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

INTERNATIONAL SEAH ... REPORT



A. CLASSIFICATION OF SUBJECT MATTER
IPC 6 C12N15/36 C12N15/49 A61K48/00

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC 6-A61K

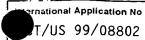
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
WO 96 36366 A (DOW STEVE W ; ELMSLIE ROBYN E (US); NAT JEWISH CENTER FOR IMMUNOLO) 21 November 1996 (1996-11-21)	1,2,7,8, 10-21, 27-29
page 6, Tine 19 -page 8, Tine 18 page 17, line 24 -page 20, line 3 page 53, line 4 -page 56, line 23 page 89, line 5 -page 92, line 15	3-6, 23-26
WO 94 28916 A (BRITISH BIOTECH PHARM; COMER MICHAEL BERISFORD (GB); MCCOURT MATTH) 22 December 1994 (1994-12-22) page 4, line 31 -page 5, line 11	1,2,5-8, 10-21, 25-29 3,4,23,
page 14, line 24 -page 17, line 27 page 25, line 20 -page 26, line 6	24
	WO 96 36366 A (DOW STEVE W ;ELMSLIE ROBYN E (US); NAT JEWISH CENTER FOR IMMUNOLO) 21 November 1996 (1996-11-21) page 6, line 19 -page 8, line 18 page 17, line 24 -page 20, line 3 page 53, line 4 -page 56, line 23 page 89, line 5 -page 92, line 15 WO 94 28916 A (BRITISH BIOTECH PHARM; COMER MICHAEL BERISFORD (GB); MCCOURT MATTH) 22 December 1994 (1994-12-22) page 4, line 31 -page 5, line 11 page 14, line 24 -page 17, line 27 page 25, line 20 -page 26, line 6

X Further documents are listed in the continuation of box C.	χ Patent family members are listed in annex.			
° Special categories of cited documents :				
"A" document defining the general state of the art which is not considered to be of particular relevance	 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to 			
"E" earlier document but published on or after the international filing date				
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another	involve an inventive step when the document is taken alone			
citation or other special reason (as specified)	"Y" document of particular relevance; the claimed invention			
"O" document referring to an oral disclosure, use, exhibition or other means	cannot be considered to involve an inventive step when the document is combined with one or more other such docu— ments, such combination being obvious to a person skilled in the art.			
"P" document published prior to the international filing date but				
later than the priority date claimed	"&" document member of the same patent family			
Date of the actual completion of the international search	Date of mailing of the international search report			
19 October 1999	03/11/1999			
Name and mailing address of the ISA	Authorized officer			
European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,				
Fax: (+31-70) 340-3016	Sitch, W			

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C.(Continu	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
	Shallon of document, with indication, where appropriate, of the relevant passages	nelevant to claim No.
X	WO 98 15285 A (WISTAR INST ;ERTL HILDEGUND C J (US); THURIN MAGDALENA (US)) 16 April 1998 (1998-04-16) page 4, line 24 -page 6, line 28	1,7, 10-20, 27-29
X	WO 96 11279 A (US HEALTH) 18 April 1996 (1996-04-18)	1,2,5-7, 10-21, 25-29
	page 18, line 20 -page 21, line 29 page 24, line 29 -page 28, line 7	
Y	KHUDYAKOV Y E ET AL: "LINEAR B-CELL EPITOPES OF THE NS3-NS4-NS5 PROTEINS OF THE HEPATITISC VIRUS AS MODELED WITH SYNTHETIC PEPTIDES" VIROLOGY, vol. 206, 1 January 1995 (1995-01-01), pages 666-672, XP000574456 ISSN: 0042-6822 page 666 abstract	3,4,23,
Y	MOLDOVEANU Z ET AL: "IMMUNE RESPONSES INDUCED BY ADMINISTRATION OF ENCAPSIDATED POLIOVIRUS REPLICONS WHICH EXPRESS HIV-1 GAG AND ENVELOPE PROTEINS" VACCINE, vol. 13, no. 11, 1 August 1995 (1995-08-01), pages 1013-1022, XP000571592 ISSN: 0264-410X page 1013 abstract	5,6,25, 26
A	WO 97 19696 A (LUSSO PAOLO ;GALLO ROBERT C (US); COCCHI FIORENZA (US); VICO ANTHO) 5 June 1997 (1997-06-05) page 4, line 11 -page 5, line 20	
A	DILLOO ET AL: "COMBINED CHEMOKINE AND CYTOKINE GENE TRANSFER ENHANCES ANTITUMOR IMMUNITY" NATURE MEDICINE, vol. 2, no. 10, October 1996 (1996-10), pages 1090-1095, XP002119425 page 1090 abstract	
E	WO 99 29728 A (GALLO ROBERT C ;DEVICO ANTHONY L (US); GARZINO DEMO ALFREDO (US);) 17 June 1999 (1999-06-17) page 12, paragraph 3 -page 25, paragraph 2	1,2,5,6, 8-22, 25-29

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mation on patent family members

ternational Application No.

Patent document cited in search report		Publication date	Patent family member(s)	Publication date
WO 9636366	A	21-11-1996	US 5705151 A US 5935568 A AU 704012 B AU 5801696 A CA 2221305 A EP 0850071 A JP 11508762 T	06-01-1998 10-08-1999 01-04-1999 29-11-1996 21-11-1996 01-07-1998 03-08-1999
WO 9428916	Α	22-12-1994	AU 6974294 A EP 0703784 A JP 8511263 T US 5925568 A ZA 9404258 A	03-01-1995 03-04-1996 26-11-1996 20-07-1999 15-12-1995
W0 9815285	Α	16-04-1998	AU 4907797 A	05-05-1998
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WO 9719696	Α	05-06-1997	AU 1141997 A EP 0869812 A	19-06-1997 14-10-1998
WO 9929728	Α	17-06-1999	AU 1815899 A	28-06-1999





WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



I C I International Dureau

INTERNATIONAL APPLICATION PUBLISH	HED U	INDER THE PATENT COOPERATION TREATY (PCT)
(51) International Patent Classification ⁶ :		(11) International Publication Number: WO 99/53960
C12N 15/36, 15/49, A61K 48/00	A3	(43) International Publication Date: 28 October 1999 (28.10.99)
(21) International Application Number: PCT/US (22) International Filing Date: 22 April 1999 (CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
(30) Priority Data: 60/082,600 22 April 1998 (22.04.98)	τ	Published With international search report. Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.
(63) Related by Continuation (CON) or Continuation-in (CIP) to Earlier Application US Filed on 60/082, 22 April 1998	600 (C	(88) Date of publication of the international search report: 23 December 1999 (23.12.99)
(71) Applicant (for all designated States except US): CORPORATION [US/US]; Intellectual Property P.O. Box 8097, Emeryville, CA 94622-8097 (US	- K4	ON 40,
(72) Inventor; and (75) Inventor/Applicant (for US only): PALIARE [US/US]; Chiron Corporation, P.O. Box 8097, E CA 94622-8097 (US).), Xav Imeryvi	ier lle,
(74) Agents: HARBIN, Alisa, A. et al.; Chiron Control Intellectual Property - R440, P.O. Box 8097, ECA 49662-8097 (US).	orporati Emeryvi	on, lie,

(54) Title: ENHANCING IMMUNE RESPONSES TO GENETIC IMMUNIZATION BY USING A CHEMOKINE

(57) Abstract

The immune response to a DNA immunogen in a mammal can be enhanced by administration of a chemokine or a polynucleotide encoding the chemokine. This method can be used, for example, to immunize or vaccinate a mammal against an infectious disease or a tumor.

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

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DE	Germany	LI	Liechtenstein				
DK	Denmark	LK	Sri Lanka	SE	Sweden		
EE	Estonia	LR	Liberia	SG	Singapore		

A. CLASSIFICATION OF SUBJECT MATTER
IPC 6 C12N15/36 C12N15/49

A61K48/00

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 6 A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X Y	WO 96 36366 A (DOW STEVE W ;ELMSLIE ROBYN E (US); NAT JEWISH CENTER FOR IMMUNOLO) 21 November 1996 (1996-11-21) page 6, line 19 -page 8, line 18	1,2,7,8, 10-21, 27-29 3-6, 23-26
	page 17, line 24 -page 20, line 3 page 53, line 4 -page 56, line 23 page 89, line 5 -page 92, line 15	
X	WO 94 28916 A (BRITISH BIOTECH PHARM;COMER MICHAEL BERISFORD (GB); MCCOURT MATTH) 22 December 1994 (1994-12-22) page 4, line 31 -page 5, line 11	1,2,5-8, 10-21, 25-29 3,4,23,
•	page 14, line 24 -page 17, line 27 page 25, line 20 -page 26, line 6	24

Further documents are listed in the continuation of box C.	X Patent family members are listed in annex.
 Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance 	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
"E" earlier document but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed	"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. "&" document member of the same patent family
Date of the actual completion of the international search 19 October 1999	Date of mailing of the international search report 03/11/1999
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer Sitch, W

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X	WO 98 15285 A (WISTAR INST ;ERTL HILDEGUND C J (US); THURIN MAGDALENA (US)) 16 April 1998 (1998-04-16) page 4, line 24 -page 6, line 28	1,7, 10-20, 27-29
X	WO 96 11279 A (US HEALTH) 18 April 1996 (1996-04-18)	1,2,5-7, 10-21, 25-29
	page 18, line 20 -page 21, line 29 page 24, line 29 -page 28, line 7	
Υ	KHUDYAKOV Y E ET AL: "LINEAR B-CELL EPITOPES OF THE NS3-NS4-NS5 PROTEINS OF THE HEPATITISC VIRUS AS MODELED WITH SYNTHETIC PEPTIDES" VIROLOGY, vol. 206, 1 January 1995 (1995-01-01), pages 666-672, XP000574456 ISSN: 0042-6822 page 666 abstract	3,4,23,
Y	MOLDOVEANU Z ET AL: "IMMUNE RESPONSES INDUCED BY ADMINISTRATION OF ENCAPSIDATED POLIOVIRUS REPLICONS WHICH EXPRESS HIV-1 GAG AND ENVELOPE PROTEINS" VACCINE, vol. 13, no. 11, 1 August 1995 (1995-08-01), pages 1013-1022, XP000571592 ISSN: 0264-410X page 1013 abstract	5,6,25, 26
A	WO 97 19696 A (LUSSO PAOLO ;GALLO ROBERT C (US); COCCHI FIORENZA (US); VICO ANTHO) 5 June 1997 (1997-06-05) page 4, line 11 -page 5, line 20	
Α	DILLOO ET AL: "COMBINED CHEMOKINE AND CYTOKINE GENE TRANSFER ENHANCES ANTITUMOR IMMUNITY" NATURE MEDICINE, vol. 2, no. 10, October 1996 (1996-10), pages 1090-1095, XP002119425 page 1090 abstract	
E	WO 99 29728 A (GALLO ROBERT C ;DEVICO ANTHONY L (US); GARZINO DEMO ALFREDO (US);) 17 June 1999 (1999-06-17) page 12, paragraph 3 -page 25, paragraph 2	1,2,5,6, 8-22, 25-29

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ational application No.

PCT/US 99/08802

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely: Remark: Although claims 11-29 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

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